

LIST OF REFERENCES CITED BY APPLICANT

(Use several sheets if necessary)

ATTY DOCKET NO. 11068-014-999	application no 10/612,604
APPLICANT Huang et al.	
FILING DATE Tuly 1 2003	GROUP 1648

			U.S. P	ATENT DOCUMENTS			
*EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
SBe	C01	5,631,128	5/97	Kozal et al.			
1	C02	5,650,268	7/97	Kozal et al.			
	C03	5,917,033	6/99	Modak et al.			
	C04	6,124,327	9/00	Silverman		_	
	C05	6,653,081	11/03	Whitcomb		_	
	C06	20040067487	4/04	Whitcomb			

			FOREIG	N PATENT DOCUMENTS				
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Spe	C07	WO99/67427	6/99	PCT				<u> </u>
_ _	C08	WO99/61658	12/99	PCT				<u> </u>
	C09	WO02/22781	9/01	PCT			0	
	C10	International Search Report of PCT/ US99/14486	6/99	PCT				
	CII	International Search Report of PCT/US01/28736	5/02	PCT	_	_		
	C12	International Search Report of PCT/US99/11629	9/99	PCT	_			
	C13	International Search Report of PCT/US01/18882	10/01	PCT				
\downarrow	C14	Copy of International Search Report PCT/US03/21024	5/04	PCT				

	OTHER REFERENCES (Including Author, Title, Date, Pertinent Pages, Etc.)					
24	<u>2</u> e	C15	Ahluwalia, G. S., et al. (1996) "2', 3'-Didehydro-3 ¹ -deoxythymidine: Regulation of its Metabolic Activation by Modulators of Thymidine-5'-triphosphate Biosynthesis" Mol. Pharm. 50: 160-165			
1		C16	Appelt, et al., 1991 "Design of Enzyme Inhibitors Using Iterative Protein Crystallographic Analysis," J. Med. Chem. 34: 1925-1934			
		C17	Arnold E., et al. (1995) "Structures of DNA and RNA Polymerases and Their Interactions with Nucleic Acid Substrates", Curr Opin Struct Biol 5:27-38;			
	-	C18	Back, KT, et al, (1996) "Reduce Replication of 3TC-Resistant HIV-1 Variants in Primary Cells Due to a Processivity Defect of the Reverse Transcriptase Enzyme", EMBO 15: 4040-4049			
V	,	C19	Balzarini J, (1998) "A Novel Mutation (F227L) Arises in the Reverse Transcriptase of Human Immunodeficiency Virus Type 1 on Dose-Escalating Treatment of HIV Type 1-Infected Cell Cultures With the Nonnucleoside Reverse Trascriptase Inhibitor Thiocarboxanilide UC-781" AIDS Res. Human, 14(3):255-260			

Be	C20	Balzarini J, et al. (1997) "Zidovudine-Resistant Human Presence Immunodeficiency Virus Type 1 Strains Subcultured in the of Both Lamivudine and Quinoxaline HBY 097 Retain Marked Sensitivity to HBY 097 but not to Lamivudine" J of Infect Dis, 176:1392-1397
	C21	Balzarini J., et al., (1992) "HIV-1-Specific Reverse Transcriptase Inhibitors Show Differential Activity Against HIV1 Mutant Strains Containing Different Amino Acid Substitutions in the Reverse Transcriptase", Virology 192:246-253
	C22	Barnes WM, (1994) "PCRAmplication of up to 35-kb DNA with High Fidelity and High Yeild from I. Bacteriophage Templates" PNAS 91:2216-2220
	C23	Bartenschlager R, et al, (1994) "Kinetic and Structural Analyses of Hepatitis C Virus Polyprotein Processing", J Virol. 68:5045-5055
	C24	Boucher CAB, et al, (1993) "High-Level Resistance to (-) Enantiomeric 2'-Deoxy-3'-Thiacytidine In Vitro is Due to One Amino Acid Substitution in the Catalytic Site of Human Immunodeficiency Virus Type 1 Reverse Transcriptase" Antimicrob Agents Chemother, 37:2231-2234
	C25	Boucher CAB, et al, (1990) "Zidovudine sensitivity of human immunodeficiency viruses from high-risk, symptom-free individuals during therapy", Lancet 336:585-590
	C26	Boyer, et al., "Analysis of Nonnucleoside Drug-Resistant Variants of Human Immunodeficiency Virus Type I Reverse Transcriptase." J. Virol., 67(4):2412-2420 (1993).
	C27	Cheeseman S.H., et al. (1995) "Phase I/II Evaluation of Defic Nevirapine Alone and in Combination with Zidvudine for Infection with Human Immunodeficiency Virus", J Acquir Immune Syndr 8:141-151
	C28	Coffin JM, (1995) "HIV Population Dynamics in Vivo: Implications for Genetic Variation, Pathogenesis, and Therapy". Science 267:483-489
	C29	Craig C and Moyle G, (1997) "The development of resistance of HIV-1 to zalcitabine", AIDS 11:271-279.
	C30	Croteau G. et al (1997) "Impaired Fitness of Human Immunodeficiency Virus Type 1 Variants with High-Leve Resistance to Protease Inhibitors" J Virol 71:1089-1096
	C31	D'Aquilla R.T. (1994) "Molecular Pathogenesis and Laboratory Monitoring", Clin Lab Med 14:393-423
	C32	De Clercq E, (1997) "Development of Resistance of Human the 4); Immunodeficiency Virus (HIV) to Anti-HI Agents: How to Prevent Problem" Intnl of Antimicro Agnts, 9:21-36
	C33	De Clerq E, (1992) "HIV Inhibitors Targeted at the Revest Transcriptase", AIDS Res. Hum Retrovin.8:119-134
	C34	DeJong, M.D., et al. (1994) "Alternating Nevirapine and Infected Zidovudine Treatment of Human Immunodeficiency Virus Type 1Persons Does Not Prolong Nevirapine Activity", J Infect Dis 169:1346-1350
	C35	DeJong MD, et al, (1996) "Host-parasite Dynamics and Outgrowth of Virus Containing a Single K7OR Amino Acid Change in Reverse Transcriptase are Responsible for the Loss of Human Immunodeficiency Virus Type RNA Load Suppression by Zidovudine", PNAS 93:5501-5506
	C36	Descamps et al., 1997 "Susceptibility of Human Immunodeficiency Virus Type 1 Group 0 Isolates to Antiretroviral Agents: In Vitro Phenotypic and Genotypic Analyses," Journal of Virology 71(11): 8893-8898.
	C37	De Antoni A., et al. Mutations in a pol gene of human immunodeficiency virus type 1 in infected patients receiving didanosine and hydroxyurea combination therapy. J. Infect Dis. (1997) OCT.; 176(4): 899-903
1	C38	Doyon L, et al, (1996) "Second Locus Involved in Human Immunodeficiency Virus Type 1 Resistance to Protease Inhibitors", J Virol 70:3763-3769
	C39	Dueweke, T.J., et al. (1993) "A Mutation in Reverse to Other Transcriptase of Bis (Heteroaryl) Piperzine Resistant Human Immunodeficiency Virus Type 1. That Confers Increased Sensitivity; Nonnucleoside Inhibitors". PNAS 90:4713-4717
	C40	Eastman, P. Scott, et al. (1995) Monisotopic Hybridization Assay for Determination of Relative Amounts of Genotypic Human Micro, Immunodeficiency Virus Type 1 Zidovudine Resistance", J Clin 2777-2780
	C41	Fitzgibbon et al. Human Immunodeficiency virus type 1 pol gene mutations in an AIDS pateint treated with multiple antiretroviral drugs. Journal of Virology, vol. 67, No. 12 (1993) pp. 7271-7275.
	C42	Frenkel et al. Specific, sensitive, and rapid assay for human immunodeficiency virus type 1 pol mutations associated with resistance to zidovudine and didanosine. Journal of Clinical Immunology. vol. 33, No. 2 (1995 pp. 342-347.
	C43	Frost, S.D.W., and McLean, A.R. (1994) "Quasispecies Dynamics and the Emergence of Drug Resistance Dur. Zidovudine Therapy of Hiv Infection", AIDS 8:323-332.
	C44	Gerondelis P, et al, (1999) "The P236L Delavirdine-Resistant Human Immunodeficiency Virus Type 1 Mutant Replication Defective and Demonstrates Alternations in both RNA 5'-End-and DNA 3',-End-Directed RNase Activities". J V irol 73: 5803-5813
	C45	Goulden MG, et al, (1996) "Selection In Vitro of an HIV-1 Variant Resistant to Both Lamivudine (3TC) and Zidvudine", AIDS 10:101-102.
$\sqrt{}$	C46	Gu Z, et al, (1994) "Identification of Novel Mutations that Confer Drug Resistance In the Human Immunodeficiency Virus Polymerase Gene", Leukemia 8(1):166-169.

SBC	C47	Hammond, et al., 1998 "Mutations in Retroviral Genes Associated with Drug Resistance," 36-79.
	C48	Harrigan PR, et al, (1998) "Relative Republication Fitness of Zidovudine-Resistant Human Immunodeficiency Virus Type 1 Isolates In Vitro", J Virol. 72:3773-3778
	C49	Ho DD, et al, (1994) "Characterization of Human Immunodeficiency Virus Type 1 Variants with Increased Resistance to a C2-Symmetric Protease Inhibitor", J Virol 68:2016-2020
	C50	Holodniy, Mark, et al. (1995) "Determination of Human Immunodefiency Virus RNA In Plasma and Cellular Viral DNA Genotypic Zidovudine Resistance Combination Therapy", J Virol, 3510-3516
	C51	Hazuda, et al., 2000 "Inhibitors of Strand Transfer That Prevent Integration and Inhibit HIV-1 Replication in Cells," Science 287: 646-650.
	C52	Iversen et al. "Multidrug-resistant immunodeficiency virus type 1 strains resulting from combination antiretroviral therapy," Journal of Virology. vol. 70, No. 2 (1996) pp. 1086-1090.
	C53	Kellam, P., et al. (1994) "Zidovudine Treatment Results in the Selection of Human Immunodeficiency Virus Type 1 Variants Whose Genotypes Confer Increasing Levels of Drug Resistance", J Gen Virol 75:341-351.
	C54	Kim EE, et al, (1995) "Crystal Structure of HIV-1 Protease in Complex with VX-478, a Potent and Orally Bioavailable Inhibitor of the Enzyme", J Am Chem Soc. 117: 1181-1182
	C55	Kleim, J., et al. (1997) "In vitro Selection for Different Mutational Patterns in the HIV-1 Reverse Transcriptase Using High and Low Selective Pressure of the Nonnucleoside Reverse Transcriptase inhibitor HBY 097" Virology. 231: 112-118
	C56	Kosalaraksa P, et al, (1999) "Comparative Fitness of Multi-Dideoxynucleoside-Resistant Human Immunodeficiency Virus Type 1 (HIV-1) in an In Vitro Competitive HIV-1 Replication Assay", J Virol 73:5356-5363.
	C57	Krebs, R., et al. 1997 "Single-Step Kinetics of HIV-1 Reverse Transcriptase Mutants Responsible for Virus Resistance to Nucleoside Inhibitors Responsible for Virus Resistance to Nucleoside Inhibitors Zidovudine and 3-TC" Biochemistry 36: 10292-10300
	C58	Kuritzkes D.R. Clinical significance of drug resistance in HIV-1 infection. AIDS (1996) vol. 10, S27-S31.
	C59	Larder BA, (1992) "3'-Azido-3'-Deoxythymidine Resistance Suppressed by a Mutation Conferring Human Immunodeficiency Virus Type 1 Resistance to Nonnucleoside Reverse Transcriptase Inhibitors", Antimicrob Agents Chemother 36: 2664-2669.
	C60	Larder BA, et al, (1991) "Zidovudine resistance predicted by direct detection of mutations in DNA from HIV-infected lymphocytes", AIDS 5:137-144.
	C61	Larder BA, et al, (1995) "Potential Mechanism for Sustained Antiretroviral Efficacy of AZT-3TC Combination Therapy", Science 269:696-699.
	C62	Lieven Stuyver, et al. (1997) "Line Probe Assay For Rapid Detection Of Drug Selected Mutations In The Human Immunodefiency Virus Type 1 Reverse Transcriptase Gene", Antimicro Aaen and Chemother, 284-291
	C63	Lin PF, et al, (1994) "Genotypic and Phenotypic Analysis of Human Immunodeficiency Virus Type 1 Isolates from Patients on Prolonged Stavudine Therapy", J Infect Disease 170:1157-1164.
	C64	Lopez-Galindez C, et al, (1991) "Characterization of genetic variation and 3'-azido-3'-deoxythymidine-resistance mutations of human immunodeficiency virus by the Rnase A mismatch cleavage method", PNAS 88:4280-4284 (Exhibit 36).
	C65	Mammamo F, et al, (1998) "Resistance-Associated Loss of Viral Fitness in Human Immunodeficiency Virus Type 1: Phenotypis Analysis of Protease and gag Coevoluation inProtease Inhibitor-Treated Patients", J Wrol 72:7632-7637
	C66	Maschera B, et al, (1996) "Mutations in the Viral Protease that Confer Resistance to Saquinavir Increase the Dissociation Rate Constant of the Protease-Saquinavir Complex",) Bio Chem 271:33231-33235.
	C67	Mayers DL, et al, (1992) "Characterization of HIV Isolates Arising After Prolonged Zidovudine Therapy", J Acq Imm Def Synd 5:749-759
	C68	Moyle GJ (1996) "Use of Viral Resistance Patterns to Antiretroviral Drugs in Optimising Selection of Drug Combinations and Sequences", Drugs 52:168-185
	C69	Mohri, H., et al. (1993) "Quantitation of Zidovudine Resistant Human Immunodeficiency Virus Type 1 in the Blood of Treated and Untreated Patients", PNAS 90:25-29
	C70	Mulligan RC and Berg P, (1980) "Expression of a Bacterial Gene in Mammalian Cells", Science 209:1422-11427
	C71	Nájera, I., et al. (1994) "Natural Occurrence of Drug Resistance Mutations in the Reverse Transcriptase of Human Immunodeficiency Virus Type 1 Isolates", Aids Res Hum Retroviruses 10:1479-1488
	C72	Nájera, I., et al. (1995) "pol Gene Quasispecies of Human Immunodeficiency Virus: Mutations Associated with Drug Resistance in virus from Patients Undergoing No Drug Therapy", <u>J Virol</u> 69:23-31
\bigvee	C73	Nunberg, J.H., et al. (1990) "Viral Resistance to Human Immunodeficiency Virus Type 1-Specific Pyridinone Reverse Transcriptase Inhibitors", J Virol 65:4887-4892

	C74	Pelemans H, et al. (1997) "Characteristics of the Pro225His Mutation in Human Immunodeficiency Virus Type 1
Spe		(HIV-1) Reverse Transcriptase That Appears Under Selective Pressure of Dose Escalating Quinoxaline Treatment of HIV-1" J. Viro, 71(11):8195-8203
	C75	Richman, D.D. et al. (1994) "Nevirapine Resistance Mutations of Human Immunodeficiency Virus Type 1 Selected during Therapy", J Virol 68:1660-1666
	C76	Richman, D.D. et al. (1991) "Human Immunodeficiency Virus Type 1 Mutants Resistant to Nonnucleoside Inhibitors of Reverse Transcriptase Arise in Tissue Culture", PNAS 88:11241-11245
	C77	Sanger, et al. (1977) "DNA Sequencing with Chain-terminating Inhibitors", PNAS 88: 11241-245.
	C78	Sakar, G. and Sommer, S.S. (1990) "The "Megaprimer" Method of Site-Directed Mutagenesis" Biotech, 8(4):404-407
	C79	Shafer RW, et al, (1994) "Combination Therapy with Zidovudine and Didanosine Selects for Drug-Resistant Human Immunodeficiency Virus Type 1 Strains with Unique Patterns of pol Gene Mutations", J Infect Disease 169:722-729
	C80	Shirasaka T, et al, (1995) "Emergence of Human Immunodeficiency Virus Type 1 Variants with Resistance to Multiple Deoxynucleosides in Patients Receiving Therapy with Dideoxynucleosides", PNAS 92:2398-2402
	C81	Southern, et al. (1982) "Transformation of Mammalian Cells to Antibotic Resistance with a Bacterial Gene Under Control of the SV40 Early Region Promoter", Appl. Genet 1:327-341
	C82	Strair RK, et al. (1993) "Recombinant Retroviral Systems For the Analysis of Drug Resistant HIV" Nucl Acds Res, 21(20): 4836-4842
	C83	Sugden B, et al, (1985)"A Vector that Replicates as a Plasmid and can be Efficiently Selected in B-Lymphoblasts Transformed by Epstein-Barr Virus", Mol Cell Bio 5:410-413.
	C84	Tisdale M, et al, (1993) "Rapid In Vitro Selection of Human Immunodeficiency Virus Type 1 Resistant to 3'- Thiacytidine Inhibitors due to a Mutation in the YMDD Region of Reverse Transcriptase", PNAS 90:5653-5656.
	C85	Vacca JP, et al, (1994) "L-735,524: An Orally Bioavailable Human Immunodeficiency Virus Type 1 Protease Inhibitor", PNAS 91:4096-4100.
	C86	Villahermosa, ML, et al. "Evaluations of mixtures of wild-type HIV-1 and HIV-1 with resistance point mutations against reverse transcriptase inhibitors" Antiviral Ther. (1998); 3(4):221-227
	C87	Zennou V, (1998) "Loss of Viral Fitness Associated with Multiple Gag and Gag-Pol Processing Defects in Human Immunodeficiency Virus Type 1 Variants Selected for Resistance to Protease Inhibitors In Vivo", J. Virol., 72:3300-3306.
	C88	Zhang Y, et al, (1997) "Drug Resistance During Indinavir Therapy is Caused by Mutations in the Protease Gene and in its Gag Substrate Cleavage Sites", J Virol 71:6662-6670.
	C89	Zhang D, et al, (1994) "Resistance to 2',3'-Dideoxycytidine Conferred by a Mutation in Codon 65 of the Human Immunodeficiency Virus Type 1 Reverse Transcriptase", Antimicrob Agents Chemother 38:282-287

EXAMINER Addy B. Chen Date considered September 14, 2004	EXAMINER	Stacy B.	Chen		DATE CONSIDERED	September	14, 2004
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^{*}EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.



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Huang et al. FILING DATE	GROUP
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			U.S. P.	ATENT DOCUMENTS			
*EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
SBe	A01	5,436,131	7/25/95	Condra et al.		-	
}:	A02	5,837,464	11/17/98	Capon et al.		-	
	A03	6,103,462	8/15/00	Paulous et al.		_	
V	A04	6,242,187	6/5/01	Capon et al.		_	

			FOREIGN	PATENT	DOCUMENTS				
		DOCUMENT NUMBER	DATE		COUNTRY	CLASS	SUBCLASS	TRANSL	ATION
								YES	NO
Stac	A05	WO99/67427	6/99	PCT		_			

Seci	A06	Gervaix, et al., "A New Reporter Cell Line to Monitor HIV Infection and Drug Susceptibility in Vitro", Proc. Natl. Acad. Sci. USA (1997), Vol. 94 pgs 4653-4658.
1	A07	Herrmann, et al., "A Working Hypotheses-Virus Resistance Development As An Indicator of Specific Antiviral Activity", Ann. NY Acad Sciences (1997), 284: 632-637.
	A08	Hertogs, et al., "A Rapid Method for Simultaneous Detection of Phenotypic Resistance to Inhibitors of Protease and Réverse Transcriptase in Recombinant Human Immunodeficiency Virus Type 1 Isolates From Patients Treated with Antiretroviral Drugs", Antimicrobial Agents and Chemotherapy (1998) 42(2): 269-276.
	A09	Lie, et al., "Advances In Quantitative PCR Technology: 5' Nuclease Assays", Curr Opinion Biotech (1998), 9(1): 43, 48.
	A10	Nijhuis, et al., "Implications of Antiretroviral Resistance on Viral Fitness", Curr. Opin. Infect Diseases (2001), 14: 28.
	A11	Petropoulos, et al., "A Novel Phenotypic Drug Susceptibility Assay For Human Immunodeficiency Virus Type 1", Antimicrobial Agents and Chemotherapy (2000), 44(4): 920-928.
	A12	Race, et al., "Analysis of HIV Cross-Resistance to Protease Inhibitors Using A Rapid Single-Cycle Recombinant Virus Assay For Patients Failing On Combination Therapies", AIDS (1999), 13(15): 2061-2068.
V	A13	Shi, et al., "A Recombinant Retroviral System for Rapid In Vivo Analysis of Human Immunodeficiency Virus Type Susceptibility to reverse Transcriptase Inhibitors", Antimicrobial Agents and Chemotherapy (1997) 41(12): 2781-85

EXAMINER Staly B. Chen DATE CONSIDERED September 14, 200
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U.S. PATENT DOCUMENTS							
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SBe	B01	6,033,902	3/7/00	Haseltine et al.			
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OTHER REFERENCES (Including Author, Title, Date, Pertinent Pages, Etc.)					
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